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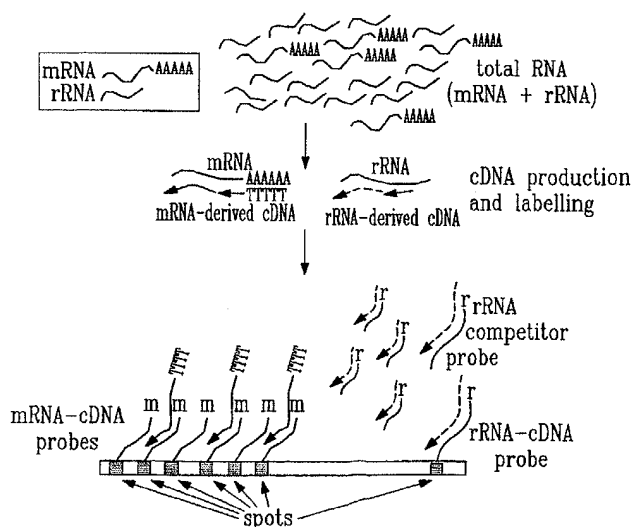
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[Continued on next page]

(54) Title: METHOD FOR NORMALIZING THE RELATIVE INTENSITIES OF DETECTION SIGNALS IN HYBRIDIZATION ARRAYS



(57) Abstract: The present invention relates to rRNA-derived cDNA used as an internal standard or control to achieve normalization of hybridization signal detection in microarray biochip technology. Because of its relatively invariant expression across tissues and treatments, 18S and 28S ribosomal RNAs are ideal internal controls for quantitative RNA analysis. A way to circumvent the technical difficulties of using ribosomal RNA as a control, because of its overabundance relative to that of other RNAs, is described and claimed in the present application. Improved methods, arrays and kits comprising arrays and free unlabelled ribosomal probes, are objects of this invention. The unlabelled ribosomal probes are used to compete out the excess or ribosomal nucleics present in a sample wherein all cDNA species of the sample are labelled before being placed in contact with the arrays.



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According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

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Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

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## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	<p>SCHEMA M ET AL: "QUANTITATIVE MONITORING OF GENE EXPRESSION PATTERNS WITH A COMPLEMENTARY DNAMICROARRAY" SCIENCE, AMERICAN ASSOCIATION FOR THE ADVANCEMENT OF SCIENCE,, US, vol. 270, no. 5235, 20 October 1995 (1995-10-20), pages 467-470, XP000644675 ISSN: 0036-8075 page 467, paragraph 4 -page 469, paragraph 2</p> <p style="text-align: center;">--- -/--</p>	1-3

☒ Further documents are listed in the continuation of box C.

☒ Patent family members are listed in annex.

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C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	EICKHOFF B ET AL: "NORMALIZATION OF ARRAY HYBRIDIZATION EXPERIMENTS IN DIFFERENTIAL GENE EXPRESSION ANALYSIS" NUCLEIC ACIDS RESEARCH, IRL PRESS LTD., OXFORD, GB, vol. 27, no. 22, 15 November 1999 (1999-11-15), pages E33I-E33III, XP001018017 ISSN: 0305-1048 page E331 -----	1-3
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**INTERNATIONAL SEARCH REPORT**  
Information on patent family members

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